PROSPECTUS



Up to 10,460,586 Shares of Common Stock Offered by the Selling Stockholders

PYXIS ONCOLOGY, INC.

This prospectus relates to the resale, from time to time by the selling stockholders named in this prospectus (the "selling stockholders") of up to 10,460,586 shares of our common stock, which consist of (i) 8,849,371 shares of our common stock held by the selling stockholders (the "Initial Shares"), and (ii) 1,611,215 shares of our common stock (the "Pre-Funded Warrant Shares") issuable upon the exercise of pre-funded warrants (held by the selling stockholders (the "Pre-Funded Warrants"). The Initial Shares and Pre-Funded Warrant Shares shall be collectively referred to as the "Securities" or the "Shares."

Our registration of shares of common stock covered by this prospectus does not mean that the selling stockholders will offer or sell any such shares. The selling stockholders received the Initial Shares and Pre-Funded Warrants from us pursuant to a private placement transaction, which was consummated on February 29, 2024. We are registering the offer and resale of the Securities to satisfy a covenant set forth in the registration rights agreement entered into on February 26, 2024 executed concurrently with a securities purchase agreement as of the same date with respect to the private placement, pursuant to which we agreed to register the resale of the Securities within a limited period of time following the date of the registration rights agreement.

We will not receive any of the proceeds from the sale of our common stock by the selling stockholders, although we will receive proceeds from the nominal exercise price of any Pre-Funded Warrants.

Any shares of our common stock subject to resale hereunder will have been issued by us and received by the selling stockholders prior to any resale of such shares pursuant to this prospectus.

The selling stockholders, or their donees, pledgees, transferees or other successors-in-interest may offer or resell the Shares from time to time through public or private transactions at prevailing market prices, at prices related to prevailing market prices or at privately negotiated prices. The selling stockholders will bear all commissions and discounts and similar selling expenses, if any, attributable to the sale of Shares. We will bear all costs, expenses and fees (other than commissions and discounts and similar selling expenses) in connection with the registration of the Shares. For additional information on the methods of sale that may be used by the selling stockholders, see "Plan of Distribution" beginning on page 9 of this prospectus.

Our common stock is listed on the Nasdaq Global Select Market under the symbol "PYXS." On March 26, 2024, the last reported sale price of our common stock was \$3.83.

We are an "emerging growth company" and a "smaller reporting company" under federal securities laws and as such, have elected to comply with reduced public company reporting requirements for this prospectus and the documents incorporated by reference herein and may elect to comply with reduced public company reporting requirements in future filings. See "Summary — Implications of Being an Emerging Growth Company and Smaller Reporting Company."

Investing in our securities involves significant risks. We strongly recommend that you read carefully the risks we describe in this prospectus and in any accompanying prospectus supplement, as well as the risk factors that are incorporated by reference into this prospectus from our filings made with the Securities and Exchange Commission. See "Risk Factors" on page 5 of this prospectus.

NEITHER THE SECURITIES AND EXCHANGE COMMISSION NOR ANY STATE SECURITIES COMMISSION HAS APPROVED OR DISAPPROVED OF THESE SECURITIES OR PASSED UPON THE ADEQUACY OR ACCURACY OF THIS PROSPECTUS. ANY REPRESENTATION TO THE CONTRARY IS A CRIMINAL OFFENSE.

The date of this prospectus is April 3, 2024

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ABOUT THIS PROSPECTUS

This prospectus is part of a registration statement on Form S-3 that we filed with the Securities and Exchange Commission (the "SEC"), using a "shelf" registration process. The selling stockholders may resell, from time to time, in one or more offerings, shares of our common stock offered by this prospectus. Information about the selling stockholders may change over time. When the selling stockholders sell shares of our common stock under this prospectus, we will, if necessary and required by law, provide a prospectus supplement that will contain specific information about the terms of that offering. Any prospectus supplement may also add to, update, modify or replace information contained in this prospectus. If a prospectus supplement is provided and the description of the offering in the prospectus supplement varies from the information in this prospectus, you should rely on the information in the prospectus supplement. You should carefully read this prospectus and the accompanying prospectus supplement, if any, along with all of the information incorporated by reference herein and therein, before making an investment decision.

You should rely only on the information contained or incorporated by reference in this prospectus and any applicable prospectus supplement. We have not, and the selling stockholders have not, authorized any other person to provide you with different or additional information. If anyone provides you with different or additional information, you should not rely on it. This prospectus is not an offer to sell, nor are the selling stockholders seeking an offer to buy, the shares offered by this prospectus in any jurisdiction where the offer and sale is not permitted. No offers or sales of any of the shares of our common stock are to be made in any jurisdiction in which such an offer or sale is not permitted. You should assume that the information contained in this prospectus or any applicable prospectus supplement is accurate only as of the date on the front cover thereof or the date of the document incorporated by reference, regardless of the time of delivery of this prospectus or any applicable prospectus supplement or any sales of the shares of our common stock offered hereby or thereby.

You should read the entire prospectus and any prospectus supplement and any related issuer free writing prospectus, as well as the documents incorporated by reference into this prospectus or any prospectus supplement or any related issuer free writing prospectus, before making an investment decision. Neither the delivery of this prospectus or any prospectus supplement or any issuer free writing prospectus nor any sale made hereunder shall under any circumstances imply that the information contained or incorporated by reference herein or in any prospectus supplement or issuer free writing prospectus, as applicable. You should assume that the information appearing in this prospectus, any prospectus supplement or any document incorporated by reference herein or therein is accurate only as of the date of the applicable documents, regardless of the time of delivery of this prospectus or any sale of securities. Our business, financial condition, results of operation and prospects may have changed since that date.

The terms "Pyxis Oncology," the "Company," "our," "us" and "we," as used in this prospectus, refer to Pyxis Oncology, Inc., a Delaware corporation, and its subsidiaries unless we state otherwise or the context indicates otherwise.

SUMMARY

This summary highlights selected information from this prospectus and does not contain all of the information that you need to consider in making your investment decision. You should carefully read the entire prospectus, any applicable prospectus supplement and any related free writing prospectus, including the risks of investing in our securities discussed under the heading "Risk Factors" contained in this prospectus, any applicable prospectus supplement and any related free writing prospectus, and under similar headings in the other documents that are incorporated by reference into this prospectus. You should also carefully read the information incorporated by reference into this prospectus, including our consolidated financial statements, and the exhibits to the registration statement of which this prospectus is a part.

Company Overview

Pyxis Oncology, Inc. is a clinical stage company focused on defeating difficult-to-treat cancers. The Company is efficiently building next generation therapeutics that hold the potential for mono and combination therapies. PYX-201, an antibody-drug conjugate ("ADC") that uniquely targets Extradomain-B Fibronectin ("EDB+FN") within the tumor stroma, and PYX-106, a fully human Siglec-15-targeting antibody designed to block suppression of T-cell proliferation and function, are being evaluated in ongoing Phase 1 clinical studies in multiple types of solid tumors. Pyxis Oncology's therapeutic candidates are designed to kill tumor cells and to address the underlying pathologies created by cancer that enable its uncontrollable proliferation and immune evasion. Pyxis Oncology's ADC and immuno-oncology ("IO") programs employ novel and emerging strategies to target a broad range of solid tumors resistant to current standards of care.

Private Placement

On February 26, 2024, we entered into a securities purchase agreement (the "Securities Purchase Agreement") with certain institutional and accredited investors (each, a "Purchaser" and collectively, the "Purchasers"), pursuant to which we sold and issued to the Purchasers in a private placement (the "Private Placement Offering") an aggregate of 8,849,371 Initial Shares and Pre-Funded Warrants to purchase up to an aggregate of 1,611,215 of shares of our common stock. The Private Placement Offering closed on February 29, 2024. Aggregate gross proceeds to the Company in respect of the Private Placement Offering were approximately \$50 million, before deducting fees payable to the placement agents and other offering expenses payable by the Company.

The Pre-Funded Warrants are immediately exercisable upon issuance at an exercise price of \$0.001 per share and do not expire.

In connection with the Securities Purchase Agreement, we entered into a registration rights agreement (the "Registration Rights Agreement"). Pursuant to the Registration Rights Agreement and subject to customary conditions, the Purchasers have certain registration rights. The registration statement of which this prospectus is a part relates to the offer and resale of the Initial Shares and the Pre-Funded Warrant Shares underlying the Pre-Funded Warrants issued to the selling stockholders pursuant to the Securities Purchase Agreement to fulfill our contractual obligations under the Registration Rights Agreement.

Implications of Being an Emerging Growth Company and Smaller Reporting Company

The Jumpstart Our Business Startups Act (the "JOBS Act") was enacted in April 2012 with the intention of encouraging capital formation in the United States and reducing the regulatory burden on newly public companies that qualify as emerging growth companies. We are an "emerging growth company" within the meaning of the JOBS Act. We may take advantage of certain exemptions from various public reporting requirements, including the requirement that we provide more than two years of audited consolidated financial statements and related management's discussion and analysis of financial condition and results of operations, and that our internal controls over financial reporting be audited by our independent registered public accounting firm pursuant to Section 404 of the Sarbanes-Oxley Act of 2002 (the Sarbanes-Oxley Act). In addition, the JOBS Act provides that an "emerging growth company" can delay adopting new or revised accounting standards until those standards apply to private companies. We intend to take advantage of these exemptions until we are no longer an emerging growth company. We have elected to use the extended transition period to enable us to comply with new or revised accounting standards that have different effective dates for public and private companies until the earlier of the date we (1) are no longer an emerging growth company and (2) affirmatively and irrevocably opt out of the extended transition period provided in the JOBS Act. As a result, our consolidated financial statements may not be companies that comply with new or revised accounting pronouncements as of public company effective dates.

We will cease to be an emerging growth company upon the earliest of (1) the end of the fiscal year following the fifth anniversary of our initial public offering; (2) the last day of the fiscal year during which our annual gross revenues are \$1.235 billion or more; (3) the date on which we have, during the previous three-year period, issued more than \$1.0 billion in non-convertible debt securities; and (4) the end of any fiscal year in which the market value of our common stock held by non-affiliates exceeded \$700.0 million as of the end of the second quarter of that fiscal year.

Additionally, we are a "smaller reporting company meaning that the market value of our stock held by non-affiliates plus the proposed aggregate amount of gross proceeds to us as a result of this offering is less than \$700 million and our annual revenue was less than \$100 million during the most recently completed fiscal year. Smaller reporting companies may take advantage of certain reduced disclosure obligations, including, among other things, providing only two years of audited financial statements in our Annual Report on Form 10-K, and, similar to emerging growth companies, smaller reporting companies have reduced disclosure obligations regarding executive compensation. We will remain a smaller reporting company until the last day of the fiscal year in which (i) the market value of our common stocks held by non-affiliates exceeds \$250 million as of the end of that year's second fiscal quarter and our annual revenue exceeds \$100 million during such completed fiscal year, or (ii) the market value of our common stock held by non-affiliates exceeds \$700 million, regardless of our annual revenue, as of the end of that year's second fiscal quarter. To the extent we take advantage of such reduced disclosure obligations, it may also make comparison of our consolidated financial statements with other public companies difficult or impossible.

Corporate Information

We were incorporated in the state of Delaware on June 11, 2018 and launched with our first employee and Series A funding in July 2019. Our principal executive offices are located at 321 Harrison Avenue, Boston, Massachusetts 02118, and our telephone number is (617) 221-9059. Our website address is www.pyxisoncology.com. The information contained on or accessible through our website is not incorporated by reference into this prospectus, and you should not consider any information contained on, or that can be accessed through, our website as part of this prospectus or in deciding whether to purchase our common stock.

The Offering

Common stock offered by us

None.

Common stock offered by the selling stockholders

Up to 10,460,586 shares of our common stock, par value \$0.001 per share, which consist of (i) 8,849,371 shares of our common stock held by the selling stockholders and (ii) 1,611,215 shares of

8,849,371 shares of our common stock held by the selling stockholders and (ii) 1,611,215 shares of our common stock issuable upon the exercise of the Pre-Funded Warrants held by the selling

stockholders.

Common stock currently outstanding

58,133,375 (as of March 20, 2024)

The Pre-Funded Warrants

The Pre-Funded Warrants are immediately exercisable upon issuance at an exercise price of \$0.001

per share and do not expire.

Selling stockholders

All of the shares of our common stock are being offered by the selling stockholders. See "Selling Stockholders" on beginning on page 7 for additional information on the selling stockholders.

Use of Proceeds

We will not receive any proceeds from the sale of the shares in this offering by the selling stockholders, although we will receive proceeds from the nominal exercise price of any Pre-Funded Warrants. See "Use of Proceeds" beginning on page 6 for additional information on the

use of proceeds.

Plan of Distribution

The selling stockholders, or their pledgees, donees, transferees, distributees, beneficiaries or other successors-in-interest, may offer or sell the shares of our common stock offered under this prospectus from time to time through public or private transactions at our prevailing market prices, at prices related to prevailing market prices or at privately negotiated prices. The selling stockholders may also resell the shares of our common stock offered under this prospectus to or through underwriters, broker-dealers or agents, who may receive compensation in the form of discounts, concessions or commissions. See "Plan of Distribution" beginning on page 9 for additional information on the methods of sale that may be used by the selling stockholders.

Risk Factors

Investing in our securities involves risk. You should carefully read and consider the information beginning on page 5 of this prospectus set forth under the heading "Risk Factors" and all other information set forth in this prospectus and the documents incorporated herein and therein by

reference before deciding to invest in our common stock.

Nasdaq symbol for common stock

"PYXS"

RISK FACTORS

Investing in our securities involves risk. You should carefully consider the specific risks discussed or incorporated by reference into the applicable prospectus supplement, together with all the other information contained in the prospectus or incorporated by reference into this prospectus and the applicable prospectus supplement. You should also consider the risks, uncertainties and assumptions discussed under the caption "Risk Factors" included in our Annual Report on Form 10-K for the year ended December 31, 2023, which is incorporated by reference into this prospectus. These risk factors may be amended, supplemented or superseded from time to time by other reports we file with the SEC in the future or by a prospectus supplement relating to a particular offering of our securities. These risks and uncertainties are not the only risks and uncertainties we face. Additional risks and uncertainties not presently known to us, or that we currently view as immaterial, may also impair our business. If any of the risks or uncertainties described in our SEC filings or any prospectus supplement or any additional risks and uncertainties actually occur, our business, financial condition and results of operations could be materially and adversely affected. In that case, the trading price of our securities could decline and you might lose all or part of your investment.

The sale of a substantial number of shares of our common stock in the public market, including resale of the Securities issued or issuable to the selling stockholders, could adversely affect the prevailing market price for our common stock.

We are registering for resale up to 10,460,586 shares of our common stock consisting of Initial Shares and Pre-Funded Warrant Shares underlying the Pre-Funded Warrants issued to the selling stockholders pursuant to the Securities Purchase Agreement to fulfill our contractual obligations under the Registration Rights Agreement. Sales of substantial amounts of shares of our common stock in the public market, or the perception that such sales might occur, could adversely affect the market price of our common stock. We cannot predict if and when the selling stockholders may sell such shares in the public markets. Furthermore, in the future, we may issue additional shares of our common stock or other equity or debt securities exercisable for, or convertible into, shares of our common stock. Any such issuances could result in substantial dilution to our existing stockholders and could cause our stock price to decline.

FORWARD-LOOKING STATEMENTS

This prospectus contains and incorporates by reference "forward-looking statements" within the meaning of the federal securities laws. All statements other than statements of historical fact contained or incorporated by reference in this prospectus, including statements regarding our future results of operations and financial position, business strategy and plans and objectives of management for future operations, are "forward-looking statements" for the purposes of this prospectus. These statements involve known and unknown risks, uncertainties and other important factors that may cause our actual results, performance or achievements to be materially different from any future results, performance or achievements expressed or implied by the forward-looking statements.

Unless the context requires otherwise, references in this prospectus to "Pyxis Oncology," the "Company," "us," and "our" refer to Pyxis Oncology, Inc. and its subsidiaries. In some cases, you can identify forward-looking statements by terms such as "anticipate," "believe," "can," "continue," "could," "estimate," "expect," "intend," "likely," "may," "might," "objective," "ongoing," "plan," "potential," "predict," "project," "should," "to be," "will," "would," or the negative or plural of these words, or similar expressions or variations, although not all forward-looking statements contain these words. The forward-looking statements in this prospectus are only predictions. We have based these forward-looking statements largely on our current expectations and projections about future events and financial trends that we believe may affect our business, financial condition and results of operations. These forward-looking statements speak only as of the date of this prospectus and are subject to a number of risks, uncertainties and assumptions described in the section titled "Risk Factors" and elsewhere in this prospectus. Because forward-looking statements are inherently subject to risks and uncertainties, some of which cannot be predicted or quantified, you should not rely on these forward-looking statements as predictions of future events. The events and circumstances reflected in our forward-looking statements may not be achieved or occur and actual results could differ materially from those projected in the forward-looking statements. Some of the key factors that could cause actual results to differ from our expectations include:

- our ability to develop and advance our current or future product candidates and programs, and to successfully initiate and complete clinical trials;
- the ability of our clinical trials to demonstrate the safety, purity and potency of our product candidates and other positive results;
- the size of the market opportunity for our product candidates, including our estimates of the number of patients who suffer from the cancers we are targeting;
- our manufacturing, commercialization and marketing capabilities and strategy;
- our plans to expand our pipeline of product candidates and further develop the Flexible Antibody Conjugation Technology, or FACT Platform, and the antibody-discovery platform, or APXiMAB Platform;
- the timing or likelihood of regulatory filings and approvals for our product candidates;
- regulatory developments in the United States and Europe and other foreign jurisdictions:
- our expectations and plans to obtain funding for our operations, including from our existing and potential future collaboration and licensing agreements;
- our ability to receive milestone or royalty payments under existing or future agreements;
- our expectations regarding our ability to obtain and maintain intellectual property protection for our product candidates:
- our continued reliance on third parties to manufacture our product candidates for preclinical studies, and, in the future, to conduct clinical trials and manufacture product candidates for such clinical trials; and
- our estimates regarding expenses, future revenue, capital requirements and needs for additional financing.

In addition, statements such as "we believe" and similar statements reflect our beliefs and opinions on the relevant subject. These statements are based upon information available to us as of the date of this prospectus and, although we believe such information forms a reasonable basis for such statements, such information may be limited or incomplete, and our statements should not be read to indicate that we have conducted a thorough inquiry into, or review of, all potentially available relevant information. Furthermore, if our forward-looking statements prove to be inaccurate, the inaccuracy may be material. Factors that may cause actual results to differ materially from current expectations include, among other things, those set forth in Part I, Item 1A of our Annual Report on Form 10-K for the year ended December 31, 2023, Part II, Item 1A – "Risk Factors" and any risks contained in any other documents incorporated by reference herein. In light of the significant uncertainties in these forward-looking statements, you should not regard these statements as a representation or warranty by us or any other person that we will achieve our objectives and plans in any specified time frame, or at all. Except as required by applicable law, we do not plan to publicly update or revise any forward-looking statements contained herein, whether as a result of any new information, future events or otherwise.

USE OF PROCEEDS

We will not receive any proceeds from the sale of Initial Shares or Pre-Funded Warrant Shares by the selling stockholders.

We may receive proceeds from the nominal exercise price of the Pre-Funded Warrants which, if exercised in full, would result in gross proceeds to us of approximately \$1,611. There can be no assurance that any of the Pre-Funded Warrants will be exercised.

We intend to use the net proceeds from any exercise of the Pre-Funded Warrants for working capital and general corporate purposes.

SELLING STOCKHOLDERS

The shares of common stock being offered by the selling stockholders, consisting of Initial Shares and Pre-Funded Warrant Shares underlying the Pre-Funded Warrants, in each case issued to the selling stockholders pursuant to the Securities Purchase Agreement, are being registered hereby to fulfill our contractual obligations under the Registration Rights Agreement. See "Prospectus Summary – Private Placement."

The following table sets forth the number and percentage of shares of our common stock beneficially owned by the selling stockholders as of March 20, 2024, taking into account the number of shares that may be offered under this prospectus and the number and percentage of our common stock beneficially owned by the selling stockholders assuming all of the shares offered under this prospectus are sold. Beneficial ownership is determined in accordance with the rules of the SEC and includes voting or investment power with respect to shares of our common stock. Generally, a person "beneficially owns" shares of our common stock if the person has or shares with others the right to vote those shares or to dispose of them, or if the person has the right to acquire voting or disposition rights within 60 days.

All information contained in the table below and the footnotes thereto is based upon information provided to us by the selling stockholders. The information in the table below and the footnotes thereto regarding shares of our common stock to be beneficially owned after the offering under this prospectus assumes the sale of all shares of our common stock being offered by the selling stockholders under this prospectus. The percentage of shares of our common stock owned is based on 58,133,375 shares of our common stock issued and outstanding as of March 20, 2024. Unless otherwise indicated in the footnotes to this table, we believe that the selling stockholders have sole voting and investment power with respect to the shares of our common stock indicated as beneficially owned.

As used in this prospectus, the term "selling stockholder" includes the selling stockholders named below and any donees, pledgees, transferees or other successors-in-interest selling shares of our common stock received after the date of this prospectus from the selling stockholders as a gift, pledge, or other non-sale related transfer.

The number of shares in the column "Maximum Number of Shares of Common Stock Offered" represents all of the shares of our common stock that the selling stockholders may offer under this prospectus. The columns captioned "Ownership After Offering" assume the sale of all of the shares of our common stock offered by the selling stockholders under this prospectus and that the selling stockholder does not acquire any additional shares of our common stock before the completion of the offering under this prospectus, other than through the exercise of the Pre-Funded Warrants. However, because the selling stockholders may sell all or some of the shares offered under this prospectus from time to time, or in another permitted manner, we cannot assure you as to the actual number of shares of our common stock that will be sold by the selling stockholders or that will be held by the selling stockholders after completion of any sales. The selling stockholders may sell some, all or none of the shares of our common stock offered under this prospectus. We do not know how long the selling stockholders will hold the Initial Shares or Pre-Funded Warrants, whether any will exercise the Pre-Funded Warrants, and upon such exercise, how long such selling stockholders will hold the shares of common stock before selling them, and we currently have no agreements, arrangements or understandings with the selling stockholders regarding the sale of any of the shares of common stock.

Under the terms of the Pre-Funded Warrants, the selling stockholder that holds Pre-Funded Warrants may not exercise the Pre-Funded Warrants to the extent such exercise would cause such selling stockholder, together with its affiliates and attribution parties, to beneficially own a number of shares of common stock which would exceed 9.99% of the number of shares of our common stock outstanding following such exercise (for purposes of the denominator, immediately after giving effect to the issuance of shares of common stock to be issued upon the applicable exercise of such Pre-Funded Warrants). The columns captioned "Ownership Before Offering" do not reflect this limitation.

	Ownership Before Offering			Ownership After Offering	
	Number of Shares of Common Stock Beneficially	Percentage of Shares of Common Stock Beneficially	Maximum Number of Shares of Common	Number of Shares of Common Stock Beneficially	Percentage of Shares of Common Stock Beneficially
Selling Stockholder	Owned	Owned	Stock Offered	Owned	Owned
Deep Track Biotechnology Master Fund, Ltd. (1)	4,184,100	7.20%	4,184,100	_	*
Atlas Diversified Master Fund, Ltd. (2)	2,092,050	3.60%	2,092,050	_	*
Ridgeback Capital Investments L.P. (3)	5,956,443	9.97%	1,611,215	4,345,228	7.27%
Blue Owl Healthcare Opportunities IV Public					
Investments LP (4)	1,046,025	1.80%	1,046,025	_	*
Laurion Capital Master Fund Ltd. (5)	3,861,179	6.64%	690,376	3,170,803	5.45%
StemPoint Capital Master Fund LP (6)	482,395	*	418,410	63,985	*
Blackstone CSP-MST FMAP Fund (7)	138,926	*	138,926	_	*
Monashee Pure Alpha SPV I LP (7)	125,817	*	125,817	_	*
BEMAP Master Fund Ltd. (7)	115,771	*	115,771	_	*
Mission Pure Alpha LP (7)	37,896	*	37,896	_	*

- * Indicates beneficial ownership of less than 1% of the outstanding shares of our common stock.
- (1) Consists of 4,184,100 shares of our common stock issued in the Private Placement Offering. Deep Track Capital, LP (the "Investment Manager") serves as the investment manager to Deep Track Biotechnology Master Fund, Ltd. ("Deep Track Master Fund") and may be deemed to beneficially own such shares. Deep Track Capital GP, LLC (the "General Partner")', is the General Partner of the Investment Manager. David Kroin is the Chief Investment Officer of the Investment Manager and managing member of the General Partner and may be deemed to beneficially own such shares. The business address of the Deep Track Master Fund, the Investment Manager, the General Partner and Mr. Kroin is 200 Greenwich Avenue, 3rd Floor, Greenwich, CT 06830.
- (2) Consists of 2,092,050 shares of our common stock issued in the Private Placement Offering. Balyasny Asset Management L.P. ("BAM") serves as investment adviser to Atlas Diversified Master Fund, Ltd. ("ADMF"). Dmitry Balyasny indirectly controls the general partner of BAM and shall be deemed to have sole voting and dispositive power over all of the shares. The business address of ADMF is c/o Maples Corporate Services Limited, P.O. Box 309, Ugland House, George Town, Grand Cayman KY1-1104, Cayman Islands, and the business address of BAM and Mr. Balyasny is 444 West Lake Street, 50th Floor, Chicago, IL 60606.
- (3) Consists of (i) 4,345,228 shares of our common stock previously acquired and (ii) 1,611,215 shares underlying Pre-Funded Warrants issued to Ridgeback Capital Investments L.P. ("RCILP") in the Private Placement Offering, the exercise of which is subject to a beneficial ownership limitation of 9.99% of our outstanding shares of common stock. Ridgeback Capital Investments LLC ("RCI") is the general partner of RCILP., Pursuant to an investment agreement, Ridgeback Capital Management LLC ("RCM") maintains investment and voting power with respect to the securities held or controlled by RCI. Wayne Holman, an individual, controls RCM and may be deemed to directly or indirectly exercise power to vote or dispose of the securities issued to RCILP. The business address of RCILP is c/o RCM 348 West 14th Street, Floor 4, New York, New York 10014.
- (4) Consists of 1,046,025 shares of our common stock issued in the Private Placement Offering. Blue Owl Healthcare Opportunities Advisors LLC is the investment manager of Blue Owl Healthcare Opportunities IV Public Investments LP ("Blue Owl") and has voting and investment power over the shares held by Blue Owl. Blue Owl Healthcare Opportunities Advisors LLC exercises voting and investment power through an investment committee comprised of Kevin Raidy, Timothy Anderson, Sandip Agarwala, and Brandyn Itzkowitz who each disclaims beneficial ownership over these securities. The business address of Blue Owl is 399 Park Avenue, Floor 38, New York, New York 10022.
- Consists of (i) 3,170,803 shares of our common stock previously acquired and (i) 690,376 shares of our common stock issued in the Private Placement Offering. Laurion Capital Management LP ("LP"), the investment manager of Laurion Capital Master Fund Ltd. ("Laurion"), has voting and investment power over the securities held by Laurion Capital Master Fund Ltd. Messrs. Benjamin A. Smith and Sheehan Maduraperuma are the managing members of Laurion Capital GP LLC, which is the general partner of Laurion Capital Management LP, and each of Benjamin A. Smith and Sheehan Maduraperuma reports shared voting and dispositive power over the 3,861,179 shares held by Laurion. Each of Laurion, Laurion Capital GP LLC, Benjamin A. Smith and Sheehan Maduraperuma disclaims beneficial ownership over these securities. The business address for Laurion, Mr. Smith, and Mr. Maduraperuma is c/o the LP, 360 Madison Avenue, Suite 1900, New York NY 10017.
- (6) Consists of 418,410 shares of our common stock issued in the Private Placement Offering to StemPoint Capital Master Fund LP ("StemPoint Fund"). StemPoint Capital LP is the investment advisor to StemPoint Fund and Jefferies Strategic Investments, LLC ("JSI"), and the sub-advisor to Titan Biotech Dislocation Fund SP ("TBDF"). StemPoint Capital LP exercises voting and investment power through its respective investment management agreement for both the StemPoint Fund and JSI, who disclaims beneficial ownership of the shares held by JSI. The address of these funds and accounts is 520 Madison Avenue, 19th Floor, New York, NY 10022.
- (7) Consists of (i) 138,926 shares of our common stock issued to Blackstone CSP-MST FMAP Fund ("FMAP"); (ii) 125,817 shares of our common stock issued to Monashee Pure Alpha SPV I LP ("Pure Alpha"); (iii) 115,771 shares of our common stock issued to BEMAP Master Fund Ltd ("BEMAP"); and (iv) 37,896 shares of our common stock issued to Mission Pure Alpha LP ("Mission"), all pursuant to the Private Placement Offering. BEMAP, FMAP, Mission and Pure Alpha are managed by Monashee Investment Management LLC ("Monashee Management"). Jeff Muller is CCO of Monashee Management and has voting and investment control over Monashee Management and, accordingly, may be deemed to have beneficial ownership of the shares held by BEMAP, FMAP, Mission and Pure Alpha. Jeff Muller, however, disclaims any beneficial ownership of the shares held by these entities. The business address of BEMAP, FMAP, Mission, Pure Alpha and Mr. Muller is c/o Monashee Investment Management LLC, 75 Park Plaza, 4th Floor, Boston, MA 02116.

PLAN OF DISTRIBUTION

The selling stockholders, including their pledgees, donees, transferees, distributees, beneficiaries or other successors in interest, may from time to time offer some or all of the shares of our common stock offered under this prospectus. We will not receive any of the proceeds from the sale of the shares of our common stock offered under this prospectus by the selling stockholders. We will bear all fees and expenses incident to our obligation to register the shares of our common stock offered under this prospectus.

The selling stockholders may sell all or a portion of the shares of our common stock beneficially owned by them and offered hereby from time to time directly or through one or more underwriters, broker-dealers or agents. If the shares of our common stock are sold through underwriters or broker-dealers, the selling stockholders will be responsible for underwriting discounts or commissions or agent's commissions. The shares of our common stock may be sold on any national securities exchange or quotation service on which the securities may be listed or quoted at the time of sale, in the over-the-counter market or in transactions otherwise than on these exchanges or systems or in the over-the-counter market and in one or more transactions at fixed prices, at prevailing market prices at the time of the sale, at varying prices determined at the time of sale, or at privately negotiated prices. These sales may be effected in transactions, which may involve crosses or block transactions.

The selling stockholders may use any one or more of the following methods when disposing of shares of our common stock or interests therein:

- ordinary brokerage transactions and transactions in which the broker-dealer solicits purchasers;
- block trades in which the broker-dealer will attempt to sell shares of our common stock as agent, but may position and resell a portion of the block as principal to facilitate the transaction;
- purchases by a broker-dealer as principal and resale by the broker-dealer for its account;
- an over-the-counter distribution;
- an exchange distribution in accordance with the rules of the applicable exchange;
- privately negotiated transactions;
- short sales effected after the effective date of the registration statement of which this prospectus forms a part;
- through the writing or settlement of options or other hedging transactions, whether through an options exchange or otherwise;
- broker-dealers may agree with the selling stockholders to sell a specified number of such shares at a stipulated price per share;
- · a combination of any such methods of sale; or
- any other method permitted pursuant to applicable law.

The selling stockholders may, from time to time, pledge or grant a security interest in some or all of the shares of our common stock owned by them and, if they default in the performance of their secured obligations, the pledgees or secured parties may offer and sell the shares of our common stock, from time to time, under this prospectus, or under an amendment to this prospectus under Rule 424(b)(3) or other applicable provision of the Securities Act amending the list of selling stockholders to include the pledgee, transferee, or other successors in interest as selling stockholder under this prospectus. The selling stockholders also may transfer the shares of our common stock in other circumstances, in which case the transferees, pledgees or other successors in interest will be the selling beneficial owners for purposes of this prospectus.

In connection with the sale of shares of our common stock or interests therein, the selling stockholders may enter into hedging transactions with broker-dealers or other financial institutions, which may in turn engage in short sales of shares of our common stock in the course of hedging the positions they assume. The selling stockholders may also sell shares of our common stock short and deliver these securities to close out its short positions, or loan or pledge the shares of our common stock to broker-dealers that in turn may sell these securities. The selling stockholders may also enter into option or other transactions with broker-dealers or other financial institutions or the creation of one or more derivative securities which require the delivery to such broker-dealer or other financial institution of shares of our common stock offered under this prospectus, which shares of our common stock such broker-dealer or other financial institution may resell pursuant to this prospectus (as supplemented or amended to reflect such transaction).

Broker-dealers engaged by the selling stockholders may arrange for other broker-dealers to participate in sales. If the selling stockholders effect certain transactions by selling shares of our common stock to or through underwriters, broker-dealers or agents, such underwriters, broker-dealers or agents may receive commissions in the form of discounts, concessions or commissions from the selling stockholders or commissions from purchasers of the shares of our common stock for whom they may act as agent or to whom they may sell as principal. Such commissions will be in amounts to be negotiated, but, except as set forth in a supplement to this prospectus, in the case of an agency transaction will not be in excess of a customary brokerage commission in compliance with applicable rules of the Financial Industry Regulatory Authority, Inc. ("FINRA"); and in the case of a principal transaction a markup or markdown in compliance with applicable FINRA rules.

The aggregate proceeds to the selling stockholders from the sale of the shares of our common stock offered under this prospectus will be the purchase price of the shares of common stock less discounts or commissions, if any. The selling stockholders reserve the right to accept and, together with their agents from time to time, to reject, in whole or in part, any proposed purchase of shares of our common stock to be made directly or through agents. We will not receive any of the proceeds from the offering of common stock under this prospectus.

The selling stockholders also may resell all or a portion of the shares of our common stock offered under this prospectus in open market transactions in reliance upon Rule 144 under the Securities Act, provided that they meet the criteria and conforms to the requirements of that rule.

The selling stockholders and any underwriters, broker-dealers or agents that participate in the sale of the shares of our common stock or interests therein may be deemed to be "underwriters" within the meaning of Section 2(a)(11) of the Securities Act. Any discounts, commissions, concessions or profit they earn on any resale of the shares of our common stock may be underwriting discounts and commissions under the Securities Act. Each selling stockholder has informed us that it does not have any written or oral agreement or understanding, directly or indirectly, with any person to distribute the securities. The selling stockholders are subject to the prospectus delivery requirements of the Securities Act.

To the extent required pursuant to Rule 424(b) under the Securities Act, the shares of our common stock to be sold, the name of the selling stockholders, the purchase price and public offering price, the names of any agents, dealer or underwriter, and any applicable commissions or discounts with respect to a particular offer will be set forth in an accompanying prospectus supplement or, if appropriate, a post-effective amendment to the registration statement that includes this prospectus.

We have agreed to keep this prospectus effective until the earliest to occur of the following events: (i) the date on which the selling stockholders shall have resold or otherwise disposed of all the Securities covered hereby; and (ii) the date on which the Securities no longer constitute "Registrable Securities" as such term is defined in the Registration Rights Agreement, such that they may be resold by the selling stockholders without registration and without regard to any volume or manner-of-sale limitations and without current public information pursuant to Rule 144 as set forth in a written opinion letter to such effect, addressed, delivered and acceptable to our transfer agent and the affected holders. In order to comply with the securities laws of some states, if applicable, the shares of our common stock may be sold in these jurisdictions only through registered or licensed brokers or dealers. In addition, in some states the shares of our common stock may not be sold unless the shares been registered or qualified for sale or an exemption from registration or qualification requirements is available and is complied with.

The selling stockholders and any other person participating in a sale of shares of our common stock registered under this prospectus will be subject to applicable provisions of the Exchange Act, and the rules and regulations thereunder, including, without limitation, to the extent applicable, Regulation M of the Exchange Act, which may limit the timing of purchases and sales of any of the shares of our common stock by the selling stockholders and any other participating person. All of the foregoing may affect the marketability of the shares of our common stock and the ability of any person or entity to engage in market-making activities with respect to the shares of our common stock. In addition, we will make copies of this prospectus (as it may be supplemented or amended from time to time) available to the selling stockholders for the purpose of satisfying the prospectus delivery requirements of the Securities Act. The selling stockholders may indemnify any broker-dealer that participates in transactions involving the sale of the shares of our common stock against certain liabilities, including liabilities arising under the Securities Act.

LEGAL MATTERS

The validity of the securities being offered hereby will be passed upon for us by Sidley Austin LLP, San Francisco, California.

EXPERTS

Ernst & Young LLP, independent registered public accounting firm, has audited our consolidated financial statements included in our Annual Report on Form 10-K for the year ended December 31, 2023, as set forth in their report, which is incorporated by reference in this prospectus and elsewhere in the registration statement. Our financial statements are incorporated by reference in reliance on Ernst & Young LLP's report, given on their authority as experts in accounting and auditing.

The consolidated financial statements of Apexigen, Inc. as of December 31, 2022 and 2021, and for the years then ended, incorporated in this prospectus by reference from the Current Report on Form 8-K/A of Pyxis Oncology, Inc. dated October 27, 2023, have been audited by Moss Adams LLP, an independent registered public accounting firm, as stated in their report (which report expresses an unqualified opinion and includes an explanatory paragraph relating to a going concern uncertainty and an emphasis of a matter paragraph relating to the reverse recapitalization), which is incorporated herein by reference. Such consolidated financial statements are incorporated by reference in reliance upon the report of such firm given their authority as experts in accounting and auditing.

WHERE YOU CAN FIND MORE INFORMATION

We have filed with the SEC a registration statement on Form S-3 under the Securities Act of 1933, as amended (the "Securities Act"), with respect to the securities offered by this prospectus and any applicable prospectus supplement. This prospectus and any applicable prospectus supplement do not contain all of the information set forth in the registration statement and its exhibits and schedules in accordance with SEC rules and regulations. For further information with respect to us and the securities being offered by this prospectus and any applicable prospectus supplement, you should read the registration statement, including its exhibits and schedules. Statements contained in this prospectus and any applicable prospectus supplement, including documents that we have incorporated by reference, as to the contents of any contract or other document referred to are not necessarily complete, and, with respect to any contract or other document filed as an exhibit to the registration statement or any other such document, each such statement is qualified in all respects by reference to the corresponding exhibit. You should review the complete contract or other document to evaluate these statements. You may obtain copies of the registration statement and its exhibits via the SEC's website at http://www.sec.gov.

We file annual, quarterly and current reports, proxy statements and other documents with the SEC under the U.S. Securities Exchange Act of 1934, as amended (the "Exchange Act"). The SEC maintains a website that contains reports, proxy and information statements and other information regarding issuers, including us, that file electronically with the SEC. You may obtain documents that we file with the SEC at http://www.sec.gov. We also make these documents available on our website at www.pyxisoncology.com. Our website and the information contained or accessible through our website is not incorporated by reference in this prospectus or any prospectus supplement, and you should not consider it part of this prospectus or any prospectus supplement.

INFORMATION INCORPORATED BY REFERENCE

SEC rules permit us to incorporate information by reference in this prospectus and any applicable prospectus supplement. This means that we can disclose important information to you by referring you to another document filed separately with the SEC. The information incorporated by reference is considered to be part of this prospectus and any applicable prospectus supplement, except for information superseded by information contained in this prospectus or any applicable prospectus supplement itself or in any subsequently filed incorporated document. This prospectus and any applicable prospectus supplement incorporate by reference the documents set forth below that we have previously filed with the SEC (Commission File No. 001-39069), other than information in such documents that is deemed to be furnished and not filed. These documents contain important information about us and our business and financial condition.

- Annual Report on Form 10-K for the year ended December 31, 2023, filed with the SEC on March 21, 2024;
- Our Current Reports on Form 8-K filed on February 28, 2024, March 13, 2024 (Item 5.02 only), March 27, 2024 (filed at 7:49 a.m. Eastern Time) and March 27, 2024 filed at 4:09 p.m. Eastern Time) and on Form 8-K/A filed on October 27, 2023; and
- The description of our common stock contained in our Registration Statement on <u>Form 8-A</u>, filed with the SEC on October 5, 2021, and any other amendment or report filed for the purpose of updating such description.

All documents that we file (but not those that we furnish) pursuant to Section 13(a), 13(c), 14 or 15(d) of the Exchange Act, after the date of the initial registration statement of which this prospectus is a part and prior to the effectiveness of the registration statement shall be deemed to be incorporated by reference into this prospectus and will automatically update and supersede the information in this prospectus, and any previously filed documents. All documents that we file (but not those that we furnish) pursuant to Section 13(a), 13(c), 14 or 15(d) of the Exchange Act on or after the date of this prospectus and prior to the termination of the offering of any of the securities covered under this prospectus shall be deemed to be incorporated by reference into this prospectus and will automatically update and supersede the information in this prospectus, the applicable prospectus supplement and any previously filed documents.

Any statement contained herein or in a document incorporated or deemed to be incorporated by reference in this prospectus or any applicable prospectus supplement shall be deemed to be modified or superseded for purposes of this prospectus and such applicable prospectus supplement to the extent that a statement contained in this prospectus or such applicable prospectus supplement, or in any other subsequently filed document which also is or is deemed to be incorporated by reference in this prospectus and such applicable prospectus supplement, modifies or supersedes such earlier statement. Any statement so modified or superseded shall not be deemed, except as so modified or superseded, to constitute a part of this prospectus or such applicable prospectus supplement.

You can obtain any of the filings incorporated by reference into this prospectus or any applicable prospectus supplement through us or from the SEC through the SEC's website at http://www.sec.gov. Upon written or oral request, we will provide, without charge, a copy of any or all of the reports and documents referred to above which have been incorporated by reference into this prospectus or any applicable prospectus supplement. Prospective investors may obtain documents incorporated by reference in this prospectus or any applicable prospectus supplement by requesting them in writing or by telephone from us at our executive offices at:

Pyxis Oncology, Inc. 321 Harrison Avenue Boston, Massachusetts, 02118 (617) 221-9059

Our reports and documents incorporated by reference herein may also be found in the "Investors" section of our website at www.pyxisoncology.com. The content of our website and any information that is linked to or accessible from our website (other than our filings with the SEC that are incorporated by reference, as set forth under "Incorporation of Certain Documents by Reference") is not incorporated by reference into this prospectus or any applicable prospectus supplement and you should not consider it a part of this prospectus, any applicable prospectus supplement, or the registration statement.

Up to 10,460,586 Shares of Common Stock Offered by the Selling Stockholders



PROSPI	ECTUS

April 3, 2024

We have not authorized any dealer, salesperson or other person to give any information or represent anything not contained in this prospectus. You must not rely on any unauthorized information. If anyone provides you with different or inconsistent information, you should not rely on it. This prospectus does not offer to sell any securities in any jurisdiction where it is unlawful. Neither the delivery of this prospectus, nor any sale made hereunder, shall create any implication that the information in this prospectus is correct after the date hereof.